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| | | | | Application Number | 10/621,485 |
| | | | | Filing Date | July 16, 2003 |
| | | | | First Named Inventor | Mueckler et al. |
| | | | | Art Unit | 1614 |
| | | | | Examiner Name | Susan Emily Fernandez |
| Sheet | 1 | of | 6 | Attorney Docket Number | 56029-41936 |

| U.S. PATENT DOCUMENTS | | | | | |
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| Examiner Initials | Cite No. ¹ | Document Number | Publication Date MM-DD-YYYY | Name of Patentee or Applicant of Cited Document | Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear |
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| AA | ALESSI, D. et al., Mechanism of Activation of Protein Kinase B by Insulin and IGF-1, The EMBO Journal, 1996, 15 (23): 6541-6551 | | | | |
| AB | ALESSI, D. et al., 3-Phosphoinositide-Dependent Protein Kinase-1 (PDK1): Structural and Functional Homology with the Drosophila DSTPK61 Kinase, Current Biology, September 18, 1997, 7 (10): 776-789 | | | | |
| AC | ALESSI, D. et al., Characterization of a 3-Phosphoinositide-Dependent Protein Kinase Which Phosphorylates and Activates Protein Kinase Ba, Current Biology, March 19, 1997, 7 (4): 261-269 | | | | |
| AD | BALENDRAN, A. et al., PDK1 Acquires PDK2 Activity in the Presence of a Synthetic Peptide Derived from the Carboxyl Terminus of PRK2, Current Biology, April 8, 1999, 9 (8): 393-404, S1-S3 | | | | |
| AE | BEHN-KRAPPA and NEWTON, The Hydrophobic Phosphorylation Motif of Conventional Protein Kinase C is Regulated by Autophosphorylation, Current Biology, June 30, 1999, 9 (14): 728-737 | | | | |
| AF | BRAZIL and HEMMINGS, Ten Years of Protein Kinase B Signalling: A Hard Act to Follow, TRENDS in Biochemical Sciences, November 2001, 26 (11): 657-664 | | | | |

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| AA | AG | CALDERHEAD, D. et al., Insulin Regulation of the Two Glucose Transporters in 3T3-L1 Adipocytes, The Journal of Biological Chemistry, August 15, 1990, 285 (23): 13800-13808 | | | | |
| AA | AH | CLARK, S. et al., Intracellular Localization of Phosphatidylinositide 3-Kinase and Insulin Receptor Substrate-1 in Adipocytes: Potential Involvement of a Membrane Skeleton, The Journal of Cell Biology, March 9, 1998, 140 (5): 1211-1225 | | | | |
| AA | AI | COFFER, P. et al., Protein Kinase B (c-Akt): A Multifunctional Mediator of Phosphatidylinositol 3-Kinase Activation, Biochem. J., 1998, Great Britain, 335: 1-13 | | | | |
| AA | AJ | CLARKE, J. et al., Research Communication – Inhibition of the Translocation of GLUT1 and GLUT4 in 3T3-L1 Cells by the Phosphatidylinositol 3-Kinase Inhibitor, Wortmannin, Biochem. J., 1994, Great Britain, 300: 631-635 | | | | |
| AA | AK | CRITCHLEY, David R., Focal Adhesions – the Cytoskeletal Connection, Current Opinion in Cell Biology, 2000, 12: 133-139 | | | | |
| AA | AL | CURRIE, R. et al., Role of Phosphatidylinositol 3,4,5-Trisphosphate in Regulating the Activity and Localization of 3-Phosphoinositide-Dependent Protein Kinase-1, Biochem. J., 1999, Great Britain, 337: 575-583 | | | | |
| AA | AM | DEDHAR, S. et al., Integrin-Linked Kinase (ILK): A Regulator of Integrin and Growth-Factor Signalling, <i>trends in CELL BIOLOGY</i> , August 1999, 9: 319-323 | | | | |
| AA | AN | DELCOMMENNE, M. et al., Phosphoinositide-3-OH Kinase-Dependent Regulation of Glycogen Synthase Kinase 3 and Protein Kinase B/AKT by the Integrin-Linked Kinase, Proc. Natl. Acad. Sci. USA, Cell Biology, September 1998, 95: 11211-11216 | | | | |
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| AA | AP | EBINA, Y. et al., The Human Insulin Receptor cDNA: The Structural Basis for Hormone-Activated Transmembrane Signalling, Cell, April 1985, 40: 747-758 | | | | |
| AA | AQ | ENDEMANN, G. et al., Phosphatidylinositol Kinase or an Associated Protein is a Substrate for the Insulin Receptor Tyrosine Kinase, The Journal of Biological Chemistry, January 5, 1990, 265 (1): 396-400 | | | | |

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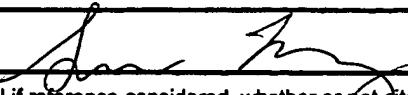
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| | | | | Examiner Name | Susan Emily Fernandez |
| Sheet | 3 | of | 6 | Attorney Docket Number | 56029-41936 |

| OTHER PRIOR ART – NON PATENT LITERATURE DOCUMENTS | | | | |
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| T ² | | | | |
| jh | AR | GAREN and LEVINTHAL, A Fine-Structure Genetic and Chemical Study of the Enzyme Alkaline Phosphatase of <i>E. Coli</i> , <i>Biochim. Biophys. Acta</i> , 1960, 38: 470-483 | | |
| jh | AS | GORDON, Julius A., Use of Vanadate as Protein-Phosphotyrosine Phosphatase Inhibitor, <i>Methods in Enzymology</i> , 1991, 201: 477-483 | | |
| jt | AT | HELLER-HARRISON, R. et al., Insulin Regulation of Membrane-Associated Insulin Receptor Substrate 1, <i>The Journal of Biological Chemistry</i> , October 13, 1995, 270 (41): 24442-24450 | | |
| jt | AU | HILL, M. et al., A Role for Protein Kinase B β /Akt2 in Insulin-Stimulated GLUT4 Translocation in Adipocytes, <i>Molecular and Cellular Biology</i> , Nov. 1999, 19 (11): 7771-7781 | | |
| jj | AV | INOUE, G. et al., Development of an <i>In Vitro</i> Reconstitution Assay for Glucose Transporter 4 Translocation, <i>PNAS</i> , December 21, 1999, 96 (26): 14919-14924 | | |
| jj | AW | JAMES, D. et al., Molecular Cloning and Characterization of an Insulin-Regulatable Glucose Transporter, <i>Nature</i> , 1989, 338: 83-87 | | |
| jt | AX | JARETT, Leonard, Subcellular Fractionation of Adipocytes, <i>Methods Enzymol</i> 31: 60-71 | | |
| jh | AY | KELLY and RUDERMAN, Insulin-Stimulated Phosphatidylinositol 3-Kinase – Association with a 185-kDa Tyrosine-Phosphorylated Protein (IRS-1) and Localization in a Low Density Membrane Vesicle, <i>The Journal of Biological Chemistry</i> , February 25, 1993, 268 (6): 4391-4398 | | |
| jh | AZ | KHWAJA, A. et al., Matrix Adhesion and Ras Transformation Both Activate a Phosphoinositide 3-OH Kinase and Protein Kinase B/Akt Cellular Survival Pathway, <i>The EMBO Journal</i> , 1997, 16 (10): 2783-2793 | | |
| jh | BA | LAVAN and LIENHARD, The Insulin-Elicited 60-kDa Phosphotyrosine Protein in Rat Adipocytes is Associated with Phosphatidylinositol 3-Kinase, <i>The Journal of Biological Chemistry</i> , March 16, 1993, 268 (8): 5921-5928 | | |
| jh | BB | LAWLOR and ALESSI, PKB/Akt: A Key Mediator of Cell Proliferation, Survival and Insulin Responses?, <i>Journal of Cell Science</i> , 2001, 114: 2903-2910 | | |

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| <i>JF</i> | BC | LYNCH, et al., Integrin-Linked Kinase Regulates Phosphorylation of Serine 473 of Protein Kinase B by an Indirect Mechanism, Oncogene, 1999, 18: 8024-8032 | | | | |
| <i>JF</i> | BD | McDONALD, J. et al., Ability of Insulin to Increase Calcium Binding by Adipocyte Plasma Membranes, Proceedings of the National Academy of Sciences of the United States of America, May 1976, 73 (5): 1542-1546 | | | | |
| <i>JF</i> | BE | PERSAD, S. et al., Regulation of Protein Kinase B/Akt-Serine 473 Phosphorylation by Integrin-Linked Kinase – Critical Roles for Kinase Activity and Amino Acids Arginine 211 and Serine 343, The Journal of Biological Chemistry, July 20, 2001, 276 (29): 27462-27469 | | | | |
| <i>JF</i> | BF | PIPER, R. et al., Differential Sorting of Two Glucose Transporters Expressed in Insulin-Sensitive Cells, Am. J. Physiol., 1991, 260 (Cell Physiol. 29): C570-C580 | | | | |
| <i>JF</i> | BG | REED, B. et al., Alterations in Insulin Binding Accompanying Differentiation of 3T3-L1 Preadipocytes, Proceedings of the National Academy of Sciences of the United States of America, Nov. 1977, 74 (11): 4876-4880 | | | | |
| <i>JF</i> | BH | RICE, K. et al., Regulation of Expression of pp160, a Putative Insulin Receptor Signal Protein, by Insulin, Dexamethasone, and 1-Methyl-3-Isobutyxanthine in 3T3-L1 Adipocytes, The Journal of Biological Chemistry, May 15, 1992, 267 (14): 10163-10167 | | | | |
| <i>JF</i> | BI | RUBIN, C. et al., Acquisition of Increased Hormone Sensitivity During <i>in Vitro</i> Adipocyte Development, The Journal of Biological Chemistry, May 25, 1977, 252 (10): 3554-3557 | | | | |
| <i>JF</i> | BJ | RUDERMAN, N. et al., Activation of Phosphatidylinositol 3-Kinase by Insulin, Proc. Natl. Acad. Sci., February 1990, 87 (Cell Biology): 1411-1415 | | | | |
| <i>JF</i> | BK | SALTIEL and KAHN, Insulin Signalling and the Regulation of Glucose and Lipid Metabolism, Nature, 2001, 414: 799-806 | | | | |
| <i>JF</i> | BL | SCHLEMMER and SIROTKA, Energy-Dependent Efflux of Methotrexate in L1210 Leukemia Cells – Evidence for the Role of an ATPase Obtained with Inside-Out Plasma Membrane Vesicles, The Journal of Biological Chemistry, July 25, 1992, 267 (21): 14746-14752. | | | | |
| <i>JF</i> | BM | SEALS, J. et al., Insulin Effect on Protein Phosphorylation of Plasma Membranes and Mitochondria in a Subcellular System from Rat Adipocytes, The Journal of Biological Chemistry, August 10, 1979, 254 (15): 6991-6996 | | | | |

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| SP | BN | SHEPHERD, P. et al., Phosphoinositide 3-Kinase: The Key Switch Mechanism in Insulin Signalling, <i>Biochem. J.</i> , 1998, Great Britain, 333: 471-490 | | | | |
| SP | BO | SIMPSON, I. et al., Insulin-Stimulated Translocation of Glucose Transporters in the Isolated Rat Adipose Cells: Characterization of Subcellular Fractions, <i>Biochimica et Biophysica Acta</i> , 1983, 763: 393-407 | | | | |
| SP | BP | SUMMERS, S. et al., Differentiation-Dependent Suppression of Platelet-Derived Growth Factor Signaling in Cultured Adipocytes, <i>The Journal of Biological Chemistry</i> , August 20, 1999, 274 (34): 23858-23867 | | | | |
| SP | BQ | TAKAKURA, K. et al., Rapid and Irreversible Inactivation of Protein Tyrosine Phosphatases PTP1B, CD45, and LAR by Peroxynitrite, <i>Archives of Biochemistry and Biophysics</i> , September 15, 1999, 369 (2): 197-207 | | | | |
| SP | BR | TOKER, A. et al., Cellular Signaling: Pivoting Around PDK-1, <i>Cell</i> , October 13, 2000, 103: 185-188 | | | | |
| SP | BS | TOKER, A. et al., Akt/Protein Kinase B is Regulated by Autophosphorylation at the Hypothetical PDK-2 Site, <i>The Journal of Biological Chemistry</i> , March 24, 2000, 275 (12): 8271-8274 | | | | |
| SP | BT | TORDJMAN, K. et al., Differential Regulation of Two Distinct Glucose Transporter Species Expressed In 3T3-L1 Adipocytes: Effect of Chronic Insulin and Tolbutamide Treatment, <i>Proceedings of the National Academy of Sciences of the United States of America</i> , October 15, 1989, 86 (20): 7761-7765 | | | | |
| SP | BU | ULLRICH, A. et al., Human Insulin Receptor and Its Relationship to the Tyrosine Kinase Family of Oncogenes, <i>Nature</i> , February 28, 1985, 313: 756-761 | | | | |
| SP | BV | VANHAESEBROECK and ALESSI, The PI3K-PDK1 Connection: More Than Just a Road to PKB, <i>Biochem. J.</i> , 2000, Great Britain, 346: 561-576 | | | | |
| SP | BW | WATSON, R. et al., Lipid Raft Microdomain Compartmentalization of TC10 is Required for Insulin Signaling and GLUT4 Translocation, <i>The Journal of Cell Biology</i> , August 20, 2001, 154 (4): 829-840 | | | | |
| SP | BX | WHITE, Morris F., The IRS-Signalling System: A Network of Docking Proteins that Mediate Insulin Action, <i>Molecular and Cellular Biochemistry</i> , 1998, 182: 3-11 | | | | |

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| AF | BY | WHITE and KAHN, The Insulin Signaling System, The Journal of Biological Chemistry, January 7, 1994, 269 (1): 1-4 | | | | |
| AF | BZ | WILLIAMS, M. et al., The Role of 3-Phosphoinositide-Dependent Protein Kinase 1 in Activating AGC Kinases Defined in Embryonic Stem Cells, Current Biology, April 5, 2000, 10 (8): 439-448 | | | | |
| AF | CA | STEPHENS, L. et al., Protein Kinase B Kinases That Mediate Phosphatidylinositol 3,4,5-Trisphosphate-Dependent Activation of Protein Kinase B, Science, January 30, 2998, 279: 710-714 | | | | |
| AF | CB | KRIAUCIUNAS, K. et al., Cellular Compartmentalization in Insulin Action: Altered Signaling by a Lipid-Modified IRS-1, Molecular and Cellular Biology, Sept. 2000, 20 (18): 6849-6859 | | | | |
| AF | CC | INOUE, G. et al., Dynamics of Insulin Signaling in 3T3-L1 Adipocytes – Differential Compartmentalization and Trafficking of Insulin Receptor Substrate (IRS)-1 and IRS-2, The Journal of Biological Chemistry, May 8, 1998, 273 (19): 11548-11555 | | | | |
| AF | CD | CROSS, D. et al., Inhibition of Glycogen Synthase Kinase-3 by Insulin Mediated by Protein Kinase B, Nature, December 1995, 378: 785-789 | | | | |
| AF | CE | HOLGADO-MADRUGA, M. et al, A Grb2-Associated Docking Protein in EGF- and Insulin-Receptor Signalling, Nature, February 1996, 379: 560-563 | | | | |
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